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    Recombinant Mycobacterium BCG adjuvant in vaccination
     Laeufer, Albrecht; Eisele, Bernd; ***Grode, Leander***
    Vakzine Projekt Management G.m.b.H., Germany
PA
SO
    Eur. Pat. Appl., 17 pp.
     CODEN: EPXXDW
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     Patent
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                      KIND DATE APPLICATION NO. DATE
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    EP 1649869
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expressing listeriolysin as adjuvant in vaccination)
IT Hemolysins
RL: BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(listeriolysins 0: ***urease*** -deficient Mycobacterium BCG
(listeriolysins 0; ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant in vaccination)
expressing listeriolysin as adjuvant in vaccination) II Antigens
expressing listeriolysin as adjuvant in vaccination) IT Antigens Tumor antigens
expressing listeriolysin as adjuvant in vaccination) II Antigens Tumor antigens RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
expressing listeriolysin as adjuvant in vaccination) IT Antigens Tumor antigens RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (microbial; ***urease*** -deficient Mycobacterium BCG expressing
expressing listeriolysin as adjuvant in vaccination) II Antigens Tumor antigens RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(non-small-cell carcinoma; ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as vaccine adjuvant for cytokine-transgenic cell immunogens) TT Lysosome (phagolysosome: ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant in vaccination in relation to) ΙT Carcinoma ***urease*** -deficient Mycobacterium BCG expressing (prostatic; listeriolysin as vaccine adjuvant for cytokine-transgenic cell immunogens) Carcinoma (pulmonary non-small-cell; ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as vaccine adjuvant for cytokine-transgenic cell immunogens) ΤТ Kidney, neoplasm ***urease*** -deficient Mycobacterium BCG (renal cell carcinoma; expressing listeriolysin as vaccine adjuvant for cytokine-transgenic cell immunogens) TТ Carcinoma (renal cell; ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as vaccine adjuvant for cytokine-transgenic cell immunogens) ΙT Head and Neck, neoplasm (squamous cell carcinoma; ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as vaccine adjuvant for cytokine-transgenic cell immunogens) Vaccines ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant for) TТ MSP-1 (protein) RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant for) Plasmodium falciparum TT (***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant for merozoite surface protein of) Malaria (***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant for vaccination against) TT Mycobacterium BCG Mycobacterium bovis (***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant in vaccination) ΤТ Antigen-presenting cell Brain, neoplasm Dendritic cell Mammary gland, neoplasm Melanoma Neoplasm (***urease*** -deficient Mycobacterium BCG expressing listeriolysin as vaccine adjuvant for cytokine-transgenic cell immunogens) TT Antimalarials Antitumor agents (vaccines: ***urease*** -deficient Mycobacterium BCG expressing

listeriolysin as adjuvant for)

IT Interferons

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(.gamma.; in combination with ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant in vaccination)

IT 884349-82-0

RL: BSU (Biological study, unclassified); PRP (Properties); THU

(Therapeutic use); BIOL (Biological study); USES (Uses)

(amino acid sequence; ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant in vaccination)

IT 9002-13-5D, ***Urease*** , subunit C

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(deficiency; ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant in vaccination)

IT 884349-81-9, DNA (Listeria monocytogenes gene hyl)

RL: BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(nucleotide sequence; ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant in vaccination)

- L2 ANSWER 2 OF 3 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 1
- AN 2005:517633 BIOSIS <<LOGINID::20080330>>

DN PREV200510303569

- TI Increased vaccine efficacy against tuberculosis of recombinant Mycobacterium bovis bacille Calmette-Guerin mutants that secrete listeriolysin.
- AU ***Grode, Leander*** ; Seiler, Peter; Baumann, Sven; Hess, Juergen; Brinkmann, Volker; Eddine, Ali Nasser; Mann, Peggy; Goosmann, Christian; Bandermann, Silke; Smith, Debbie; Bancroft, Gregory J.; Reyrat, Jean-Marc; van Soolingen, Dick; Raupach, Barbell; Kaufmann, Stefan H. E. [Reprint Author]
- CS Max Planck Inst Infect Biol, Dept Immunol, Schumannstr 21-22, D-10117 Berlin, Germany

Kaufmann@mpiib-Berlin.mpg.de

- SO Journal of Clinical Investigation, (SEP 2005) Vol. 115, No. 9, pp. 2472-2479.
- CODEN: JCINAO. ISSN: 0021-9738.
- DT Article
- LA English
- ED Entered STN: 23 Nov 2005
 - Last Updated on STN: 23 Nov 2005
- AB The tuberculosis vaccine Mycobacterium bovis bacille Calmette-Guerin (BCG) was equipped with the membrane-perforating listeriolysin (HHy) of Listeria monocytogenes, which was shown to improve protection against Mycobacterium tuberculosis. Following aerosol challenge, the HHy-secreting recombinant BCG (hHy(+) rBCG) vaccine was shown to protect significantly better against aerosol infection with M. tuberculosis than did the parental BCG strain. The isogenic, **urease** C-deficient hHy(+) rBCG (Pelta ureC hHy(+) rBCG) vaccine, providing an intraphagosomal pH closer to the acidic pH optimum for HHy activity, exhibited still higher vaccine efficacy than parental BCG. Delta ureC hHy(+) rBCG also induced profound protection against a member of the M. tuberculosis BejingoW genotype family while parental BCG failed to do so consistently. HHy not only promoted antigen translocation into the cytoplasm but also apoptosis of infected macrophages. We concluded that superior vaccine efficacy of

- Delta ureC hly(+) rBCG as compared with parental BCG is primarily based on improved cross-priming, which causes enhanced T cell-mediated immunity.
- ΑU ***Grode, Leander*** ; Seiler, Peter; Baumann, Sven; Hess, Juergen; Brinkmann, Volker; Eddine, Ali Nasser; Mann, Peggy; Goosmann, Christian; Bandermann, Silke; Smith, Debbie; . .
- AB. . . was shown to protect significantly better against aerosol infection with M. tuberculosis than did the parental BCG strain. The isogenic, ***urease*** C-deficient hlv(+) rBCG (Delta ureC hlv(+) rBCG) vaccine,
- providing an intraphagosomal pH closer to the acidic pH optimum for Hly.
- IT of Organisms
- macrophage: immune system, blood and lymphatics
- tuberculosis: bacterial disease, drug therapy
- Tuberculosis (MeSH)
- IT Chemicals & Biochemicals
 - ***urease*** [EC 3.5.1.5]; listeriolysin; bacille Calmette-Guerin: immunologic-drug, vaccine
- RN 9002-13-5 (***urease***)
- 9002-13-5 (EC 3.5.1.5)
- ANSWER 3 OF 3 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2004:927244 CAPLUS <<LOGINID::20080330>>
- DM 141:394066
- TT Vaccines comprising antigen domain and phagolysosomal escape domain for treating tuberculosis, cancer and infection
- ***Grode, Leander*** ; Kaufmann, Stefan H. E.; Raupach, Baerbel; Hess, TN Juergen
- PA Max-Planck-Gesellschaft zur Foerderung der Wissenschaften e.V., Germany SO PCT Int. Appl., 39 pp.
- CODEN: PIXXD2 Patent DT
- English T 75

FAN.		1																
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PI	WO 2004094469					2004	1104							21	0040	423		
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	EP	1618	128			A1		2006	0125		EP 2	004-	7290	90		21	0040	423
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		1798						2006									040	423
	JP	2007	5243	67		T		2007	0830		JP 2	006-	5052	50		21	0040	423

HR

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	IN	2005KN02337	A	20070727	IN	2005-KN2337	20051122
	US	2007134267	A1	20070614	US	2006-554408	20061130
I	US	2003-464644P	P	20030423			

PRAT WO 2004-EP4345 TeT 20040423

AB The present invention relates to novel recombinant vaccines comprising fusion protein contg. an antigenic domain and a phagolysosomal escape domain, providing protective immunity against tuberculosis. The antigenic domain is from Mycobacterium tuberculosis antigen Ag85B, Ag85A or ESAT-6; or Mycobacterium bovis antigen Ag85B. The antigenic domain can also be derived from autoantigen, tumor antigen, viral antigen, parasitic antigen, bacterial antigen or their immunogenic fragment. The phagolysosomal escape domain is a Listeria phagolysosomal escape domain. Further, the present invention refers to novel recombinant nucleic acid mols., vectors contg. said nucleic acid mols., cells transformed with said nucleic acid mols. and polypeptides encoded by said nucleic acid mols. These recombinant vaccines are used together with diluents, carriers and

adjuvants; and are prepd. for mucosal or parenteral administration.

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 4 ALL CITATIONS AVAILABLE IN THE RE FORMAT

TN ***Grode, Leander*** ; Kaufmann, Stefan H. E.; Raupach, Baerbel; Hess, Juergen

9002-13-5. ***Urease*** IT

RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)

(inactivation or -deficient; vaccines comprising antigen domain and phagolysosomal escape domain for treating tuberculosis, cancer and infection)

=> e kaufmann stefan h/au

E1 1 KAUFMANN STEFAN F M/AU E2 KAUFMANN STEFAN G/AU E3 4 --> KAUFMANN STEFAN H/AU E4 965 KAUFMANN STEFAN H E/AU E5 KAUFMANN STEFAN H K/AU 1 E6 KAUFMANN STEFAN HE/AU 4 E7 2 KAUFMANN STEFAN HUGO ERNST/AU E8 KAUFMANN STEFAN J E/AU 1 E9 3 KAUFMANN STEFANIE/AU E10 KAUFMANN STEFFEN/AU E11 1 KAUFMANN STEMP D/AU E12 KAUFMANN STEPHAN/AU R

=> s e3-e7 and (urease deficient)

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=> s e3-e7 and (urease)

T.3

4 ("KAUFMANN STEFAN H"/AU OR "KAUFMANN STEFAN H E"/AU OR "KAUFMANN STEFAN H K"/AU OR "KAUFMANN STEFAN HE"/AU OR "KAUFMANN STEFAN HUGO ERNST"/AU) AND (UREASE)

=> dup rem 14 PROCESSING COMPLETED FOR L4 1.5 2 DUP REM L4 (2 DUPLICATES REMOVED) => d bib ab kwic 1-YOU HAVE REQUESTED DATA FROM 2 ANSWERS - CONTINUE? Y/(N):v

- L5 ANSWER 1 OF 2 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 1
- AN 2005:517633 BIOSIS <<LOGINID::20080330>>
- DN PREV200510303569
- TI Increased vaccine efficacy against tuberculosis of recombinant Mycobacterium bovis bacille Calmette-Guerin mutants that secrete listeriolysin.
- AU Grode, Leander; Seiler, Peter; Baumann, Sven; Hess, Juergen; Brinkmann, Volker; Eddine, Ali Nasser; Mann, Peggy; Goosmann, Christian; Bandermann, Silke; Smith, Debbie; Bancroft, Gregory J.; Reyrat, Jean-Marc; van Soolingen, Dick; Raupach, Barbell; ***Kaufmann, Stefan H. E.***
 [Reprint Author]
- CS Max Planck Inst Infect Biol, Dept Immunol, Schumannstr 21-22, D-10117 Berlin, Germany Kaufmann@mpiib-Berlin.mpg.de
- SO Journal of Clinical Investigation, (SEP 2005) Vol. 115, No. 9, pp. 2472-2479.
 CODEN: JCINAO. ISSN: 0021-9738.
- DT Article
- LA English
- ED Entered STN: 23 Nov 2005
 - Last Updated on STN: 23 Nov 2005
- AB The tuberculosis vaccine Mycobacterium bovis bacille Calmette-Guerin (BCG) was equipped with the membrane-perforating listeriolysin (Hlv) of Listeria monocytogenes, which was shown to improve protection against Mycobacterium tuberculosis. Following aerosol challenge, the Hlv-secreting recombinant BCG (hly(+) rBCG) vaccine was shown to protect significantly better against aerosol infection with M. tuberculosis than did the parental BCG strain. The isogenic, ***urease*** C-deficient hly(+) rBCG (Delta ureC hly(+) rBCG) vaccine, providing an intraphagosomal pH closer to the acidic pH optimum for Hly activity, exhibited still higher vaccine efficacy than parental BCG. Delta ureC hly(+) rBCG also induced profound protection against a member of the M. tuberculosis Beijing/W genotype family while parental BCG failed to do so consistently. Hly not only promoted antigen translocation into the cytoplasm but also apoptosis of infected macrophages. We concluded that superior vaccine efficacy of Delta ureC hlv(+) rBCG as compared with parental BCG is primarily based on improved cross-priming, which causes enhanced T cell-mediated immunity.
- AU. . . Ali Nasser; Mann, Peggy; Goosmann, Christian; Bandermann, Silke; Smith, Debbie; Bancroft, Gregory J.; Reyrat, Jean-Marc; van Soolingen, Dick; Raupach, Barbell; ***Kaufmann, Stefan H. E.*** [Reprint Author] AB. . . was shown to protect significantly better against aerosol infection
- with M. tuberculosis than did the parental BCG strain. The isogenic,
 urease C-deficient hly(+) rBCG (Delta urec hly(+) rBCG) vaccine,
 providing an intraphaqosomal pH closer to the acidic pH optimum for Hly.
- of Organisms
- or organisms
 macrophage: immune system, blood and lymphatics
- IT Chemicals & Biochemicals

```
***urease*** [EC 3.5.1.5]; listeriolysin; bacille Calmette-Guerin:
       immunologic-drug, vaccine
RN
   9002-13-5 ( ***urease*** )
    9002-13-5 (EC 3.5.1.5)
```

- ANSWER 2 OF 2 CAPLUS COPYRIGHT 2008 ACS on STN L5
- ΔN 2004:927244 CAPLUS <<LOGINID::20080330>>
- DN 141:394066
- TΙ Vaccines comprising antigen domain and phagolysosomal escape domain for treating tuberculosis, cancer and infection
- IN Grode, Leander; ***Kaufmann, Stefan H. E.*** ; Raupach, Baerbel; Hess, Juergen

ADDITION NO

D3 mm

PA Max-Planck-Gesellschaft zur Foerderung der Wissenschaften e.V., Germany SO PCT Int. Appl., 39 pp.

KIND DAME

CODEN: PIXXD2 DT

DAMENIE NO

- Patent
- LA English

FAN.CNT 1

		TENT :							APPLICATION NO.											
PI					A1 2004110				WO 2004-EP4345							0040	423			
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			LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,		
			NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,		
			TJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW		
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			TD,	TG																
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	EP	1618	128					2006	0125	EP 2004-729090										
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			1798762			A 20060705			CN 2004-80010664						20040423					
		2007														20040423				
		2005														20051013				
	IN	2005	KN02:	337		A		2007	0727		IN 2	005-	KN23:	37		21	0051	122		
		2007							0614		US 2	006-	5544	8 0		21	0061	130		
PRAI		2003																		
	WO	2004	-EP4	345		W		2004	0423											

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ALL CITATIONS AVAILABLE IN THE RE FORMAT

DAUDACH D/AH

- IN Grode, Leander; ****Kaufmann, Stefan H. E.*** ; Raupach, Baerbel; Hess, Juergen
- IT 9002-13-5, ***Urease***

RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)

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=> e raupach barbel/au
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E1	3/		KAUPACH	D/AU
E2	36		RAUPACH	BAERBEL/AU
E3	22	>	RAUPACH	BARBEL/AU
E4	1		RAUPACH	BARBELL/AU
E5	7		RAUPACH	C/AU
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E8	5		RAUPACH	DALE C/AU
E9	1		RAUPACH	DALE R/AU
E10	9		RAUPACH	E/AU
E11	125		RAUPACH	F/AU
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=> s e1-e4 and urease

L6 4 ("RAUPACH B"/AU OR "RAUPACH BAERBEL"/AU OR "RAUPACH BARBEL"/AU
OR "RAUPACH BARBELL"/AU) AND UREASE

=> dup rem 16

L7

PROCESSING COMPLETED FOR L6

2 DUP REM L6 (2 DUPLICATES REMOVED)

=> d bib ab kwic 1-

YOU HAVE REQUESTED DATA FROM 2 ANSWERS - CONTINUE? Y/(N):v

- L7 ANSWER 1 OF 2 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 1
- AN 2005:517633 BIOSIS <<LOGINID::20080330>>
- DN PREV200510303569
- TI Increased vaccine efficacy against tuberculosis of recombinant Mycobacterium bovis bacille Calmette-Guerin mutants that secrete listeriolysin.
- AU Grode, Leander; Seiler, Peter; Baumann, Sven; Hess, Juergen; Brinkmann, Volker; Eddine, Ali Nasser; Mann, Peggy; Goosmann, Christian; Bandermann, Silke; Smith, Debbie; Bancroft, Gregory J.; Reyrat, Jean-Marc; van Soolingen, Dick; ***Raupach, Barbell***; Kaufmann, Stefan H. E. [Reporth Author]
- CS Max Planck Inst Infect Biol, Dept Immunol, Schumannstr 21-22, D-10117 Berlin, Germany
- Kaufmann@mpiib-Berlin.mpg.de
- SO Journal of Clinical Investigation, (SEP 2005) Vol. 115, No. 9, pp. 2472-2479.
 CODEN: JCINAO. ISSN: 0021-9738.
- DT Article

- LA English
- Entered STN: 23 Nov 2005
 - Last Updated on STN: 23 Nov 2005
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- AU. . . Volker; Eddine, Ali Nasser; Mann, Peggy; Goosmann, Christian; Bandermann, Silke; Smith, Debbie; Bancroft, Gregory J.; Revrat, Jean-Marc; van Soolingen, Dick; ***Raupach, Barbell*** ; Kaufmann, Stefan H. E. [Reprint Author]
- . . was shown to protect significantly better against aerosol infection AB. with M. tuberculosis than did the parental BCG strain. The isogenic, ***urease*** C-deficient hly(+) rBCG (Delta ureC hly(+) rBCG) vaccine, providing an intraphagosomal pH closer to the acidic pH optimum for Hly.
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macrophage: immune system, blood and lymphatics

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> tuberculosis: bacterial disease, drug therapy Tuberculosis (MeSH)

TТ

Chemicals & Biochemicals

urease [EC 3.5.1.5]; listeriolysin; bacille Calmette-Guerin: immunologic-drug, vaccine

RN 9002-13-5 (***urease***)

9002-13-5 (EC 3.5.1.5)

- ANSWER 2 OF 2 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2004:927244 CAPLUS <<LOGINID::20080330>>

DN 141:394066

- Vaccines comprising antigen domain and phagolysosomal escape domain for treating tuberculosis, cancer and infection
- TN Grode, Leander; Kaufmann, Stefan H. E.; ***Raupach, Baerbel*** ; Hess, Juergen
- Max-Planck-Gesellschaft zur Foerderung der Wissenschaften e.V., Germany SO PCT Int. Appl., 39 pp.
- CODEN: PIXXD2
- DT Patent
- LA English
- FAN.CN1

PAN.	CNI I				
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        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
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    CA 2523084
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                                                               20040423
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR
                              20060530
                                       BR 2004-9789
    BR 2004009789
                       A
                                                               20040423
    CN 1798762
                        Α
                              20060705
                                       CN 2004-80010664
                                                               20040423
    JP 2007524367
                       T
                             20070830 JP 2006-505250
                                                               20040423
    ZA 2005008276
                       A
                            20060628 ZA 2005-8276
                                                               20051013
                            20070727 IN 2005-KN2337
    IN 2005KN02337
                       A
                                                              20051122
    US 2007134267
                       A1 20070614 US 2006-554408
                                                              20061130
PRAI US 2003-464644P
                       P
                            20030423
    WO 2004-EP4345
                       W
                             20040423
```

AB The present invention relates to novel recombinant vaccines comprising fusion protein contg. an antigenic domain and a phagolysosmal escape domain. providing protective immunity against tuberculosis. The antigenic domain is from Mycobacterium tuberculosis antigen Ag83B, Ag83B or ESAT-6; or Mycobacterium bovis antigen Ag83B. The antigenic domain can also be derived from autoantigen, tumor antigen, viral antigen, parasitic antigen, bacterial antigen or their immunogenic fragment. The phagolysosomal escape domain is a Listeria phagolysosomal escape domain. Further, the present invention refers to novel recombinant nucleic acid mols. vectors contg. said nucleic acid mols., cells transformed with said nucleic acid mols. and polypeptides encoded by said nucleic acid mols. These recombinant vaccines are used together with diluents, carriers and

adjuvants; and are prepd. for mucosal or parenteral administration.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

- IN Grode, Leander; Kaufmann, Stefan H. E.; ***Raupach, Baerbel*** ; Hess, Juergen
- IT 9002-13-5, ***Urease***

RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)

(inactivation or -deficient; vaccines comprising antigen domain and phagolysosomal escape domain for treating tuberculosis, cancer and infection)

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=> e hess jurgen/au
E1
           1
              HESS JUNIOR ARTUR/AU
E2
           3
                HESS JURG/AU
E3
          38 --> HESS JURGEN/AU
E4
          2
               HESS JURGEN C/AU
E5
          2
               HESS JURGEN H/AU
          6
               HESS JUSTIN M/AU
E7
        605
               HESS K/AU
E8
         23 HESS K A/AU
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E9
           1 HESS K BELLEVILLE F/AU
2 HESS K C/AU
E10
E11
            8
                 HESS K D/AU
E12
            2.
                HESS K G/AU
=> s e2-e5 and urease
            1 ("HESS JURG"/AU OR "HESS JURGEN"/AU OR "HESS JURGEN C"/AU OR
              "HESS JURGEN H"/AU) AND UREASE
=> d
L8 ANSWER 1 OF 1
                     MEDLINE on STN
AN 2005580918
                   MEDLINE <<LOGINID::20080330>>
DN PubMed ID: 16110326
TT
    Increased vaccine efficacy against tuberculosis of recombinant
    Mycobacterium bovis bacille Calmette-Guerin mutants that secrete
     listeriolysin.
    Grode Leander; Seiler Peter; Baumann Sven; ***Hess Jurgen*** ;
     Brinkmann Volker; Nasser Eddine Ali; Mann Peggy; Goosmann Christian;
     Bandermann Silke; Smith Debbie; Bancroft Gregory J; Reyrat Jean-Marc; van
     Soolingen Dick; Raupach Barbel; Kaufmann Stefan H E
CS
    Max Planck Institute for Infection Biology, Berlin, Germany.
    The Journal of clinical investigation, (2005 Sep) Vol. 115, No. 9, pp.
SO
     2472-9. Electronic Publication: 2005-08-18.
    Journal code: 7802877. ISSN: 0021-9738.
CY
    United States
DT Journal: Article: (JOURNAL ARTICLE)
    (RESEARCH SUPPORT, NON-U.S. GOV'T)
LA English
FS
    Abridged Index Medicus Journals; Priority Journals
EM
    200512
ED Entered STN: 3 Nov 2005
    Last Updated on STN: 18 Dec 2005
     Entered Medline: 14 Dec 2005
=> s (urease deficient)
           75 (UREASE DEFICIENT)
=> dup rem 19
PROCESSING COMPLETED FOR L9
L10
            31 DUP REM L9 (44 DUPLICATES REMOVED)
=> s 110 and (bact? or mycobact? or tuberculosis or bovis)
            20 L10 AND (BACT? OR MYCOBACT? OR TUBERCULOSIS OR BOVIS)
=> d bib ab kwic 1-
YOU HAVE REQUESTED DATA FROM 20 ANSWERS - CONTINUE? Y/(N):v
L11 ANSWER 1 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
AN
    2007:355406 BIOSIS <<LOGINID::20080330>>
    PREV200700359871
    Characterization of the urease operon of Brucella abortus and assessment
    of its role in virulence of the ***bacterium***
   Sangari, Felix J.; Seoane, Asuncion; Rodriquez, Maria Cruz; Aquero, Jesus;
AU
    Garcia Lobo, Juan M. [Reprint Author]
CS Univ Cantabria, Dept Biol Mol, Fac Med, C Cardenal Herrera Oria S-N,
```

Santander 39011, Spain jmglobo@unican.es

SO Infection and Immunity, (FEB 2007) Vol. 75, No. 2, pp. 774-780. CODEN: INFIBR. ISSN: 0019-9567.

DT Article

LA English

ED Entered STN: 20 Jun 2007

Last Updated on STN: 20 Jun 2007

Most members of the genus Brucella show strong urease activity. However, ΔR the role of this enzyme in the pathogenesis of Brucella infections is poorly understood. We isolated several Tn5 insertion mutants deficient in urease activity from Brucella abortus strain 2308. The mutations of most of these mutants mapped to a 5.7-kbp DNA region essential for urease activity. Sequencing of this region, designated urel, revealed the presence of seven open reading frames corresponding to the urease structural proteins (UreA, UreB, and UreC) and the accessory proteins (UreD, UreE, UreF, and UreG). In addition to the urease genes, another gene (cobT) was identified, and inactivation of this gene affected urease activity in Brucella. Subsequent analysis of the previously described sequences of the genomes of Brucella spp. revealed the presence of a second urease cluster, ure2, in all them. The ure2 locus was apparently ***Urease*** - ***deficient*** mutants inactive in B. abortus 2308. were used to evaluate the role of urease in Brucella pathogenesis. The urease-producing strains were found to be resistant in vitro to strong acid conditions in the presence of urea, while urease-negative mutants were susceptible to acid treatment. Similarly, the urease-negative mutants were killed more efficiently than the urease-producing strains during transit through the stomach. These results suggested that urease protects brucellae during their passage through the stomach when the ***bacteria*** are acquired by the oral route, which is the major route of infection in human brucellosis.

TI Characterization of the urease operon of Brucella abortus and assessment of its role in virulence of the ***bacterium*** .

AB. . . presence of a second urease cluster, ure2, in all them. The ure2 locus was apparently inactive in B. abortus 2308. ***Urease*** - ***deficient*** mutants were used to evaluate the role of urease in Brucella pathogenesis. The urease-producing strains were found to be resistant. . . during transit through the stomach. These results suggested that urease protects brucellae during their passage through the stomach when the ***bacteria*** are acquired by the oral route, which is the major route of infection in human brucellosis.

and Assimilation); Enzymology (Biochemistry and Molecular Biophysics)

T Parts, Structures, & Systems of Organisms

stomach: digestive system

IT Diseases

brucellosis: ***bacterial*** disease, infectious disease Brucellosis (MeSH)

IT Diseases

Brucella abortus infection: ***bacterial*** disease, infectious disease

IT Chemicals & Biochemicals

DNA; urease [EC 3.5.1.5]; UreA; UreB; UreG; UreD; UreE; UreF; UreC ORGN Classifier

Gram-Negative Aerobic Rods and Cocci 06500

Super Taxa

Eubacteria; ***Bacteria*** ; Microorganisms

```
Organism Name
Brucella abortus (species): strain-2308
Taxa Notes
***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
Hominidae 86215
Super Taxa
Primates; Mammalia; Vertebrata; Chordata; Animalia
Organism Name
human (common)
Taxa Notes
Animals, . . .
```

- L11 ANSWER 2 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
- AN 2006:296770 BIOSIS <<LOGINID::20080330>>
- DN PREV200600297562
- TI The role of Klebsiella pneumoniae urease in intestinal colonization and resistance to gastrointestinal stress.
- AU Maroncle, Nathalie; Rich, Chantal; Forestier, Christiane [Reprint Author] CS Univ Auvergne, Fac Pharm, Bacteriol Lab, 28 Pl H Dunant, F-63000 Clermont Ferrand, France
 - Christiane.forestier@u-clermontI.fr
- SO Research in Microbiology, (MAR 2006) Vol. 157, No. 2, pp. 184-193. CODEN: RMCREW. ISSN: 0923-2508.
- DT Article
- LA English
- ED Entered STN: 31 May 2006
- Last Updated on STN: 31 May 2006
- AB The first step in nosocomial infections due to Klebsiella pneumoniae is colonization of the patient's gastrointestinal (GI) tract. In a previous work, signature-tagged mutagenesis was used in a murine model to identify 13 genes required for efficient colonization, two of which were involved in urea metabolism. The role of urease was further investigated by the construction and analysis of an isogenic ***urease*** -
 - ***deficient*** mutant. The behavior of both the wild-type strain and ***urease*** - ***deficient*** mutant was tested under hostile conditions, reproducing stresses encountered in the GI environment. The wild-type strain had an acid tolerance response (ATR) to inorganic acid, was resistant to organic acids (38.5% survival) and was able to survive concentrations of bile encountered in vivo. The absence of urease did not affect the resistance of K. pneumoniae to acid and bile stresses, but the enhanced adhesion response to Int-407 cells after exposure to bile observed with the wild-type strain was no longer detected with the urease mutant. When tested in the murine intestinal colonization model, both strains were mainly recovered in the large intestine parts, and the mutant was impaired in its colonization capacities, but only when tested in competition with the wild-type strain. These findings emphasize the prominent role played by metabolic function in the colonization process of such a complex ecosystem as the host GI tract. (c) 2005 Elsevier SAS. All rights reserved.
- AB. . . were involved in urea metabolism. The role of urease was further investigated by the construction and analysis of an isogenic
 - ***urease*** ***deficient*** mutant. The behavior of both the wild-type strain and the ***urease*** ***deficient*** mutant was tested under hostile conditions, reproducing stresses encountered in the GI environment. The wild-type strain had an acid tolerance. . .

```
Enterobacteriaceae 06702
     Super Taxa
        Facultatively Anaerobic Gram-Negative Rods; Eubacteria;
          ***Bacteria*** ; Microorganisms
     Organism Name
        Klebsiella pneumoniae (species): pathogen
     Taxa Notes
            ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
        Hominidae
                   86215
     Super Taxa
        Primates; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
        human (common): host
     Taxa Notes
```

- L11 ANSWER 3 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
- AN 2006:26018 BIOSIS <<LOGINID::20080330>>
- DN PREV200600025071
- Production of ammonium by Helicobacter pylori mediates occludin processing TI and disruption of tight junctions in Caco-2 cells.
- AU Lytton, Simon D. [Reprint Author]; Fischer, Wolfgang; Nagel, Wolfram; Haas, Rainer; Beck, Franz X.
- SeraDiaLogist, Hertlingstr 1, D-81545 Munich, Germany Simon.lvtton@t-online.de
- SO Microbiology (Reading), (OCT 2005) Vol. 151, No. Part 10, pp. 3267-3276. ISSN: 1350-0872.
- DT Article
- LA English ED Entered STN: 21 Dec 2005
 - Last Updated on STN: 21 Dec 2005
- Tight junctions, paracellular permeability barriers that define epithelial AR cell polarity, play an essential role in transepithelial transport, cell-cell adhesion and lymphocyte transmigration. They are also important for the maintenance of innate immune defence and intestinal antigen uptake. Ammonium (NH4+) is elevated in the gastric aspirates of Helicobacter pylori-infected patients and has been implicated in the disruption of tight-junction functional integrity and the induction of gastric mucosal damage during H. pylori infection. The precise mechanism of the effect of ammonium and the molecular targets of ammonium in host tissue are not vet identified. To study the effects of ammonium on epithelial tight junctions, the human colon carcinoma cell line Caco-2 was cultured on permeable supports and the transepithelial resistance (TER) was measured at different time intervals following exposure to ammonium salts or H. pylori-derived ammonium. A biphasic response to treatment with ammonium was found. Acute exposure to ammonium salts or NH3/NH4+ derived from urea metabolism by wild-type H. pylori resulted in a 20-30% decrease in TER. After 24 h, the NH4C1-treated cells showed a partial recovery of TER. In contrast, the control culture, or cultures that were exposed to supernatants derived from ***urease*** - ***deficient*** H. pylori, showed no significant decrease in TER. Occludin-specific immunoblots revealed the expression of a low-molecular-weight form of occludin of 42 kDa upon NH3/NH4+ exposure. The results indicate that modulation of tight-junction function by H. pylori is ammonium-dependent and linked to the accumulation of a low-molecular-weight and detergent-soluble form of occludin.
- AB. . . showed a partial recovery of TER. In contrast, the control culture,

```
or cultures that were exposed to supernatants derived from ***urease***
    - ***deficient*** H. pylori, showed no significant decrease in TER.
    Occludin-specific immunoblots revealed the expression of a
    low-molecular-weight form of occludin of. . .
ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives 06210
    Super Taxa
       Eubacteria; ***Bacteria*** ; Microorganisms
    Organism Name
       Helicobacter pylori (species): pathogen
    Taxa Notes
           ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
       Hominidae
                   86215
    Super Taxa
       Primates; Mammalia; Vertebrata; Chordata; Animalia
    Organism Name
       human (common)
       Caco-2 cell line (cell line). .
L11 ANSWER 4 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
    2004:316567 BIOSIS <<LOGINID::20080330>>
AN
DN
    PREV200400316839
    Selection and properties of Streptococcus thermophilus mutants deficient
    in urease
    Monnet, C. [Reprint Author]; Pernoud, S.; Sepulchre, A.; Fremaux, C.;
    Corrieu, G.
    Unite Mixte Rech Genie and Microbiol Proc Alimentai, INRA, F-78850,
    Thiverval Grignon, France
    monnet@grignon.inra.fr
    Journal of Dairy Science, (June 2004) Vol. 87, No. 6, pp. 1634-1640.
    print.
    CODEN: JDSCAE. ISSN: 0022-0302.
DT
   Article
LA.
    English
   Entered STN: 15 Jul 2004
    Last Updated on STN: 15 Jul 2004
AB
    Natural variations of the urea content of milk have a detrimental effect
    on the regularity of acidification by Streptococcus thermophilus strains
    used in dairy processes. The aim of the present study was to select
      ***urease*** - ***deficient*** mutants of S. thermophilus and to
    investigate their properties. Using an improved screening medium on agar
    plates, mutants were selected from 4 different parent strains after
    mutagen treatment and by spontaneous mutation. Most mutants were stable
    and had a phage sensitivity profile similar to that of their parent
    strain. Some of them contained detrimental secondary mutations, as their
    acidifying activity was lower than that of the parent strain cultivated in
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AII

type of mutant was much lower among spontaneous mutants than among mutants ***deficient*** mutants in dairy processes may have several advantages, such as an increase in acidification, an improved regularity of acidification, and a lower production of ammonia in whey.

the presence of the urease inhibitor flurofamide. The proportion of this

AB. . regularity of acidification by Streptococcus thermophilus strains used in dairy processes. The aim of the present study was to select ***urease*** - ***deficient*** mutants of S. thermophilus and to investigate their properties. Using an improved screening medium on agar

selected after mutagen treatment. Utilization of ***urease*** -

```
plates, mutants were selected. . . of this type of mutant was much
    lower among spontaneous mutants than among mutants selected after mutagen
    treatment. Utilization of ***urease*** - ***deficient*** mutants in
    dairy processes may have several advantages, such as an increase in
    acidification, an improved regularity of acidification, and. . .
ORGN Classifier
       Gram-Positive Cocci 07700
    Super Taxa
       Eubacteria:
                   ***Bacteria*** ; Microorganisms
    Organism Name
       Streptococcus thermophilus (species): ***urease*** -
         ***deficient*** mutants
    Taxa Notes
            ***Bacteria*** , Eubacteria, Microorganisms
L11 ANSWER 5 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
    2004:104625 BIOSIS <<LOGINID::20080330>>
    PREV200400096230
                  ***urease*** - ***deficient*** derivatives of
    Motility of
    Helicobacter pylori.
    Tan, Shumin; Berg, Douglas E. [Reprint Author]
    Department of Molecular Microbiology, Washington University School of
    Medicine, Campus Box 8230, St. Louis, MO, 63110, USA
    berg@borcim.wustl.edu
    Journal of Bacteriology, (February 2004) Vol. 186, No. 3, pp. 885-888.
    CODEN: JOBAAY. ISSN: 0021-9193.
    Article
   English
   Entered STN: 18 Feb 2004
    Last Updated on STN: 18 Feb 2004
AB Early studies of a ureB mutant derivative of Helicobacter pylori had
    suggested that urease is needed for motility and that urease action helps
    energize flagellar rotation. Here we report experiments showing that
    motility is unaffected by deletion of ureA and ureB (urease genes) or by
    inactivation of ureB alone, especially if H. pylori strains used as
    recipients for transformation with mutant alleles are preselected for
    motility. This result was obtained with the strain used in the early
    studies (CPY3401) and also with 15 other strains, 3 of which can colonize
    mice. We conclude that urease is not needed for H. pylori motility.
    Motility of ***urease*** - ***deficient*** derivatives of
    Helicobacter pylori.
ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives
    Super Taxa
       Eubacteria;
                   ***Bacteria*** ; Microorganisms
    Organism Name
       Helicobacter pylori (species): pathogen, motility, strain-88-3887,
       strain-A28-1, strain-A66-1, strain-CYP3401, strain-Chen13, strain-F28,
       strain-GS5, strain-HK192, strain-PCM4, strain-PeCan28, strain-R64,
       strain-R66, strain-R76, strain-R82, strain-SS1, strain-X47,
          ***urease***
                      - ***deficient***
                                          derivatives
    Taxa Notes
            ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
       Muridae 86375
    Super Taxa
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AN

DN TI

ΑU

CS

SO

DT LA

ED

```
Rodentia; Mammalia; Vertebrata; Chordata; Animalia
    Organism Name
       mouse (common)
    Taxa Notes
       Animals, . .
L11 ANSWER 6 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
    2000:418052 BIOSIS <<LOGINID::20080330>>
AN
DN
    PREV200000418052
    Dual functions of Streptococcus salivarius urease.
    Chen, Yi-Ywan M.; Weaver, Cheryl A.; Burne, Robert A. [Reprint author]
    Center for Oral Biology, University of Rochester Medical Center, 601
    Elmwood Ave., Rochester, NY, 14642, USA
    Journal of Bacteriology, (August, 2000) Vol. 182, No. 16, pp. 4667-4669.
    print.
    CODEN: JOBAAY. ISSN: 0021-9193.
    Article
LA
   English
ED
   Entered STN: 4 Oct 2000
    Last Updated on STN: 8 Jan 2002
        ***urease*** - ***deficient*** derivative of Streptococcus
AB
    salivarius 57.I was constructed by allelic exchange at the ureC locus.
    The wild-type strain was protected against acid killing through hydrolysis
    of physiologically relevant concentrations of urea, whereas the mutant was
    not. Also, S. salivarius could use urea as a source of nitrogen for
    growth exclusively through a urease-dependent pathway.
    A ***urease*** - ***deficient*** derivative of Streptococcus
    salivarius 57.I was constructed by allelic exchange at the ureC locus.
    The wild-type strain was protected against. . .
ORGN Classifier
       Gram-Positive Cocci 07700
    Super Taxa
       Eubacteria; ***Bacteria*** ; Microorganisms
    Organism Name
       Streptococcus salivarius: strain-57.I
    Taxa Notes
           ***Bacteria*** , Eubacteria, Microorganisms
L11 ANSWER 7 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
AN
    2000:400579 BIOSIS <<LOGINID::20080330>>
DN
    PREV200000400579
    Helicobacter pylori urease suppresses
                                           ***bactericidal*** activity of
    peroxynitrite via carbon dioxide production.
    Kuwahara, Hideo; Miyamoto, Yoichi; Akaike, Takaaki [Reprint author];
    Kubota, Tatsuo; Sawa, Tomohiro; Okamoto, Shinichiro; Maeda, Hiroshi
    [Reprint author]
    Department of Microbiology, Kumamoto University School of Medicine, 2-2-1
    Honjo, Kumamoto, 860-0811, Japan
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TT

CS

TI

SO

DT

LA

ED

print.

Article

English

CODEN: INFIBR. ISSN: 0019-9567.

Entered STN: 20 Sep 2000

Last Updated on STN: 8 Jan 2002 AB Helicobacter pylori can produce a persistent infection in the human stomach, where chronic and active inflammation, including the infiltration

Infection and Immunity, (August, 2000) Vol. 68, No. 8, pp. 4378-4383.

of phagocytes such as neutrophils and monocytes, is induced. H. pylori may have a defense system against the antimicrobial actions of phagocytes. We studied the defense mechanism of H. pylori against host-derived peroxynitrite (ONOO-), a ****bactericidal*** metabolite of nitric oxide, focusing on the role of H. pylori urease, which produces CO2 and NH3 from urea and is known to be an essential factor for colonization. The viability of H. pylori decreased in a time-dependent manner with continuous exposure to 1 muM ONOO-, i.e., 0.2% of the initial ***bacteria*** remained after a 5-min treatment without urea. The ***bactericidal*** action of ONOO- against H. pylori was significantly attenuated by the addition of 10 mM urea, the substrate for urease, whereas ONOO-induced killing of a ***burease*** - ***deficient*** mutant of H. pylori or Campylobacter jejuni, another microaerophilic ***bacterium**** lacking urease, was not affected by the addition of

whereas ONOO—induced killing of a ***urease*** - ***deficient***
mutant of H. pylori or Campylobacter jejuni, another microaerophilic
bacterium lacking urease, was not affected by the addition of
urea. Such as protective effect of urea was potentiated by
supplementation with exogenous urease, and it was almost completely
nullified by 10 muM flurofamide, a specific inhibitor of urease. The
bactericidal action of ONOO— was also suppressed by the addition
of 20 mM NAMPOO2 but not but by addition of 20 mM NAMPOO to the addition of 20 mM NAMPOO to the but by addition of 20 mM NAMPOO to the second of the addition of 20 mM NAMPOO to the second of the addition of 20 mM NAMPOO to the second of the addition of 20 mM NAMPOO to the second of the addition of of the addition

bactericidal action of ONDO- was also suppressed by the addition of 20 mM NABCO3 but not by the addition of 20 mM NABCO3 but not by the addition of 20 mM NABCO3.

significantly reduced by the addition of urea or NABCO3, as assessed by high-performance liquid chromatography with electrochemical detection. These results suggest that H. pylori-associated urease functions to produce a potent ONDO- scavenger, COZ/HCO3-, that defends the ****Detarial***

 $$\star\star\star$$ bacteria $\!\!\star\star\star$ from ONOO- cytotoxicity. The protective effect of urease

may thus facilitate sustained ***bacterial*** colonization in the infected gastric mucosa.

I Helicobacter pylori urease suppresses ***bactericidal*** activity of peroxynitrite via carbon dioxide production.

. . system against the antimicrobial actions of phagocytes. We studied the defense mechanism of H. pylori against host-derived peroxynitrite (ONOO-), a ***bactericidal*** metabolite of nitric oxide, focusing on the role of H. pylori urease, which produces CO2 and NH3 from urea and. . . of H. pylori decreased in a time-dependent manner with continuous exposure to 1 muM ONOO-, i.e., 0.2% of the initial ***bacteria*** remained after a 5-min treatment without urea. The ***bactericidal*** action of ONOO- against H. pylori was significantly attenuated by the addition of 10 mM urea, the substrate for urease, whereas ONOO--induced killing of a ***urease*** - ***deficient*** mutant of H. pylori or Campylobacter jejuni, another microaerophilic ***bacterium*** lacking urease, was not affected by the addition of urea. Such as protective effect of urea was potentiated by supplementation with exogenous urease, and it was almost completely nullified by 10 muM flurofamide, a specific inhibitor of urease. The ***bactericidal*** action of ONOO- was also suppressed by the addition of 20 mM NaHCO3 but not by the addition of 20. . . electrochemical detection. These results suggest that H. pylori-associated urease functions to produce a potent ONOO- scavenger, CO2/HCO3-, that defends the ***bacteria*** from ONOO- cytotoxicity. The protective effect of urease may thus facilitate sustained

IT Organisms

gastric mucosa: digestive system, infection; phagocytes: immune system; stomach: digestive system

bacterial colonization in the infected gastric mucosa.

IT Chemicals & Biochemicals carbon dioxide: production; peroxynitrite: ***bactericidal***

```
activity, nitric oxide ***bactericidal*** metabolite; urease:
        Helicobacter pylori
ORGN Classifier
        Aerobic Helical or Vibrioid Gram-Negatives 06210
     Super Taxa
        Eubacteria; ***Bacteria*** ; Microorganisms
     Organism Name
        Campylobacter jejuni: pathogen
        Helicobacter pylori: defense mechanism, pathogen
     Taxa Notes
            ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
       Hominidae 86215
     Super Taxa
        Primates; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
       human
     Taxa Notes
        Animals, Chordates, . . .
L11 ANSWER 8 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
    1999:114559 BIOSIS <<LOGINID::20080330>>
AN
DN
     PREV199900114559
TI
    Genetic and physiologic characterization of urease of Actinomyces
    naeslundii.
AII
    Morou-Bermudez, Evangelia; Burne, Robert A. [Reprint author]
CS
     Cent. Oral Biol., Univ. Rochester Med. Cent., 601 Elmwood Ave., Rochester,
    NY 14642, USA
SO
    Infection and Immunity, (Feb., 1999) Vol. 67, No. 2, pp. 504-512. print.
     CODEN: INFIBR. ISSN: 0019-9567.
    Article
T.A
    English
ED
    Entered STN: 12 Mar 1999
     Last Updated on STN: 12 Mar 1999
AB
    Ammonia production from urea by ureolytic oral ***bacteria*** is
     believed to have a significant impact on oral health and the ecological
     balance of oral microbial populations. In this study we cloned and
     characterized the urease gene cluster of Actinomyces naeslundii, which is
     one of the pioneer organisms in the oral cavity and a significant
     constituent of supragingival and subgingival dental plaque in children and
     adults. An internal fragment of the ureC gene of A. naeslundii WVU45 was
     initially amplified by PCR with degenerate primers derived from conserved
     amino acid sequences of the large catalytic subunit of urease in
       ***bacteria*** and plants. The PCR product was then used as a probe to
     identify recombinant ***bacteriophages*** carrying the A. naeslundii
     urease gene cluster and roughly 30 kbp of flanking DNA. Nucleotide
     sequence analysis demonstrated that the gene cluster was comprised of
     seven contiguously arranged open reading frames with significant
     homologies at the protein and nucleotide sequence levels to the ureABCEFGD
     genes from other organisms. By using primer extension, a putative
     transcription initiation site was mapped at 66 bases 5' to the start codon
     of ureA. A ***urease*** - ***deficient*** strain was constructed by
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insertion of a kanamycin resistance determinant within the ureC gene via allelic replacement. In contrast to the wild-type organism, the isogenic mutant was unable to grow in a semidefined medium supplemented with urea as the nitrogen source and was not protected by the addition of urea against killing in moderately acidic environments. These data indicated

that urea can be effectively utilized as a nitrogen source by A. naeslundii via a urease-dependent pathway and that ureolysis can protect A. naeslundii against environmental acidification at physiologically relevant pH values. Therefore, urease could confer to A. naeslundii critical selective advantages over nonureolytic organisms in dental plaque, constituting an important determinant of plaque ecology. Ammonia production from urea by ureolytic oral ***bacteria*** believed to have a significant impact on oral health and the ecological balance of oral microbial populations. In this. . . amplified by PCR with degenerate primers derived from conserved amino acid sequences of the large catalytic subunit of urease in ***bacteria*** and plants. The PCR product was then used as a probe to identify recombinant ***bacteriophages*** carrying the A. naeslundii urease gene cluster and roughly 30 kbp of flanking DNA. Nucleotide sequence analysis demonstrated that the. . . primer extension, a putative transcription initiation site was mapped at 66 bases 5' to the start codon of ureA. A ***urease*** - ***deficient*** strain was constructed by insertion of a kanamycin resistance determinant within the ureC gene via allelic replacement. In contrast to. . Major Concepts Enzymology (Biochemistry and Molecular Biophysics); Infection IΤ Diseases dental plaque: ***bacterial*** disease, dental and oral disease Dental Plague (MeSH) TТ Chemicals & Biochemicals ammonia: production; urea; urease; Actinomyces naeslundii ureA gene; Actinomyces. . ORGN Classifier Irregular Nonsporing Gram-Positive Rods 08890 Super Taxa Actinomycetes and Related Organisms; Eubacteria; ***Bacteria*** ; Microorganisms Organism Name Actinomyces naeslundii: pathogen, strain-WVU45 Taxa Notes ***Bacteria*** , Eubacteria, Microorganisms L11 ANSWER 9 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN AN 1996:361212 BIOSIS <<LOGINID::20080330>> DN PREV199699083568 TT Factors affecting growth and antibiotic susceptibility of Helicobacter pylori: Effect of pH and urea on the survival of a wild-type strain and a ***urease*** - ***deficient*** mutant. Sjostrom, J. E. [Reprint author]; Larsson, H. Dep. Cell Biol., Astra Hassle AB, Molndal, Sweden CS SO Journal of Medical Microbiology, (1996) Vol. 44, No. 6, pp. 425-433. CODEN: JMMIAV. ISSN: 0022-2615. DT Article LA English ED Entered STN: 14 Aug 1996 Last Updated on STN: 15 Aug 1996 AB This study investigated how pH and the presence of urea affect the survival and growth of Helicobacter pylori and whether these factors affect susceptibility to antibiotics in vitro. The viability of a wild-type strain and a ***urease*** - ***deficient*** mutant of H. pylori was studied after incubation for 1 h in buffers at different pH values at 37 degree C under microaerophilic conditions. Viable counts

were not affected at pH 5 and pH 7. In buffer at pH 3, there were no viable organisms, but urea (6.25 mm) protected the wild-type strain, which survived well. At pH 9, urea further reduced the viability of H. pylori and flurofamide almost abolished the effect of urea on the wild-type strain. Neither urea nor flurofamide affected the viability of the ***urease*** - ***deficient*** mutant under the same conditions. Growth was also pH dependent and was enhanced in shake-cultures. At pH 5, urea supported growth of the wild-type strain, but at pH 7 a toxic effect on the ***bacteria*** was observed. Growth of H. pylori at pH 5.9 was poor, and susceptibility to amoxycillin, erythromycin and clarithromycin was markedly less than at pH 7.2 and 7.9. The ***bactericidal*** activities of metronidazole and tetracycline were similar at the different pH values studied. At neutral pH the killing rates of amoxycillin and clarithromycin were growth rate dependent. Susceptibility to metronidazole was enhanced in stationary cultures. The interaction obtained between the proton pump inhibitor, omeprazole, and amoxycillin at pH 7 was of additive type. These results suggest that pH and growth conditions may be important in the antibacterial efficacy of different antibiotics in vivo and also provide a possible explanation for the potentiating effect of omeprazole with antibiotics in the treatment of H. pylori infections.

. . and antibiotic susceptibility of Helicobacter pylori: Effect of pH TI. and urea on the survival of a wild-type strain and a ***urease*** -***deficient*** mutant.

. Helicobacter pylori and whether these factors affect susceptibility

to antibiotics in vitro. The viability of a wild-type strain and a ***urease*** - ***deficient*** mutant of H. pylori was studied after incubation for 1 h in buffers at different pH values at 37 degree. . . flurofamide almost abolished the effect of urea on the wild-type strain. Neither urea nor flurofamide affected the viability of the ***urease*** ***deficient*** mutant under the same conditions. Growth was also pH dependent and was enhanced in shake-cultures. At pH 5, urea supported growth of the wild-type strain, but at pH 7 a toxic effect on the ***bacteria*** was observed. Growth of H. pylori at pH 5.9 was poor, and susceptibility to amoxycillin, erythromycin and clarithromycin was markedly less than at pH 7.2 and 7.9. The ***bactericidal*** activities of metronidazole and tetracycline were similar at the different pH values studied. At neutral pH the killing rates of. . . ORGN Classifier Aerobic Helical or Vibrioid Gram-Negatives Super Taxa

Eubacteria; ***Bacteria*** ; Microorganisms

Organism Name

aerobic helical or vibrioid gram-negative ***bacteria*** Helicobacter pylori Taxa Notes

ORGN Classifier

Bacteria , Eubacteria, Microorganisms

Hominidae 86215

AB.

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia Organism Name

human

Taxa Notes

Animals, Chordates, . . .

SIN

- AN 1996:183710 BIOSIS <<LOGINID::20080330>>
- DN PREV199698739839
- TI In vitro antibacterial activity of omeprazole and its selectivity for Helicobacter spp. are dependent on incubation conditions.
- AU Sjostrom, J. E. [Reprint author]; Fryklund, J.; Kuhler, T.; Larsson, H.
- CS Astra Hassle AB, Dep. Cell Biol., S-431 83 Molndal, Sweden
- SO Antimicrobial Agents and Chemotherapy, (1996) Vol. 40, No. 3, pp. 621-626. CODEN: AMACCO, ISSN: 0066-4804.
- DT Article
- LA English
- ED Entered STN: 29 Apr 1996
 - Last Updated on STN: 29 Apr 1996
- AB Factors affecting the in vitro antibacterial activity of omeprazole were studied. Our data show that 3H-labeled omeprazole covalently bound to Helicobacter pylori and to other gram-negative and gram-positive
 - ***bacteria*** . The compound was found to bind to a broad range of proteins in H. pylori, and at pH 5, binding was enhanced 15-fold compared with binding at pH 7. The ***bactericidal*** activity correlated to the degree of binding, and at pH 5, a pH at which omeprazole readily converts to the active sulfenamide form, beta-mercaptoethanol, a known scavenger of sulfenamide, and fetal calf serum, to which noncovalent protein binding of omeprazole is known to occur, reduced the level of binding and almost entirely abolished the ***bactericidal*** activity. At pH 7 the killing activities of omeprazole and structural analogs (e.g., proton pump inhibitors) were dependent on the time-dependent conversion (half-life) to the corresponding sulfenamide. The ***bactericidal*** activity exerted by the sulfenamide form at pH 5 was not specific for the genus Helicobacter. However, in brucella broth at pH 7 with 10% fetal calf serum, only Helicobacter spp. were susceptible to omeprazole, with MBCs in the range of 32 to 64 mu-q/ml, and MBCs for more stable proton pump inhibitors were higher. Wild-type H. pylori and its isogenic
 - ***ureage*** ***deficient*** mutant were equally susceptible to omeprazole. Thus, the urease is not a lethal target for omeprazole action in H. pylori. In conclusion, the antibacterial activities of omeprazole and analogs are dependent on pH and the composition of the medium used. Thus, at a low pH in buffer, these compounds have a nonselective action, whereas in broth at neutral pH, the mechanism of action is selective for Helicobacter spp.
- AB. . . omeprazole were studied. Our data show that 3H-labeled omeprazole covalently bound to Helicobacter pylori and to other gram-negative and gram-positive ***bacteria*** . The compound was found to bind to a broad range of proteins in H. pylori, and at pH 5, binding was enhanced 15-fold compared with binding at pH 7. The ***bactericidal*** activity correlated to the degree of binding, and at pH 5, a pH at which omeprazole readily converts to the. . . which noncovalent protein binding of omeprazole is known to occur, reduced the level of binding and almost entirely abolished the ***bactericidal*** activity. At pH 7 the killing activities of omeprazole and structural analogs (e.g., proton pump inhibitors) were dependent on the time-dependent conversion (half-life) to the corresponding sulfenamide. The ***bactericidal*** activity exerted by the sulfenamide form at pH 5 was not specific for the genus Helicobacter. However, in brucella broth. . . 32 to 64 mu-g/ml, and MBCs for more stable proton pump inhibitors were higher. Wild-type H. pylori and its isogenic ***urease*** - ***deficient*** mutant were equally susceptible to omeprazole. Thus, the urease is not a lethal target for omeprazole action in H. pylori.. . .

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ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives 06210
    Super Taxa
       Eubacteria; ***Bacteria*** ; Microorganisms
    Organism Name
       aerobic helical or vibrioid gram-negative ***bacteria***
       Helicobacter pylori
    Taxa Notes
           ***Bacteria*** , Eubacteria, Microorganisms
L11 ANSWER 11 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on
AN
    1995:315763 BIOSIS <<LOGINID::20080330>>
    PREV199598330063
DN
ΤT
    Avirulent, ***urease*** - ***deficient*** Helicobacter pylori
    colonizes gastric epithelial explants ex vivo.
    Eaton, K. A. [Reprint author]; Krakowka, S.
CS Dep. Vet. Pathobiol., OSU, 1925 Coffey Rd., Columbus, OH 43210, USA
SO Scandinavian Journal of Gastroenterology, (1995) Vol. 30, No. 5, pp.
    CODEN: SJGRA4. ISSN: 0036-5521.
DT
    Article
LA
    English
    Entered STN: 30 Jul 1995
    Last Updated on STN: 30 Jul 1995
AB
    Background: Urease-negative Helicobacter pylori generated by insertional
    mutagenesis fails to colonize gnotobiotic piglets, and this effect is
    largely independent of gastric pH. The purpose of this study was to
    determine whether urease-negative H. pylori colonizes gastric explants ex
    vivo. Methods: Gastric mucosal explants derived from neonatal germ-free
    piglets were inoculated with either wild-type H. pylori or one of two
    mutants derived by insertional mutagenesis. Results: All three
      ***bacterial*** strains colonized explants. The level of colonization
    increased over the duration of the experiment, reaching 10-8-10-9 cfu/g
    gastric mucosa by 72 h after inoculation. Morphologic evidence of
    colonization was similar to that observed in gnotobiotic piglets.
    Conclusions: Colonization of explants was not affected by lack of urease.
    These results contrast with previous findings showing that urease activity
    is essential for colonization of piglets by H. pylori. Thus,
    urease-dependent colonization is dependent on an intact gastric
    microenvironment.
                ***urease*** - ***deficient***
    Avirulent,
                                                  Helicobacter pylori
    colonizes gastric epithelial explants ex vivo.
   . . piglets were inoculated with either wild-type H. pylori or one of
    two mutants derived by insertional mutagenesis. Results: All three
      ***bacterial*** strains colonized explants. The level of colonization
    increased over the duration of the experiment, reaching 10-8-10-9 cfu/g
    gastric mucosa by. .
ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives 06210
    Super Taxa
       Eubacteria; ***Bacteria*** ; Microorganisms
    Organism Name
       aerobic helical or vibrioid gram-negative ***bacteria***
       Helicobacter pylori
    Taxa Notes
           ***Bacteria*** , Eubacteria, Microorganisms
```

```
ORGN Classifier
Suidae 85740
Super Taxa
Artiodactyla; Mammalia; Vertebrata; Chordata; Animalia
Organism Name
pig
Taxa Notes
Animals, Artiodactyls, . . .
```

- L11 ANSWER 12 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
- AN 1994:446969 BIOSIS <<LOGINID::20080330>>
- DN PREV199497459969
- TI Effect of gastric pH on urease-dependent colonization of gnotobiotic piglets by Helicobacter pylori.
- AU Eaton, Kathryn A. [Reprint author]; Krakowka, Steven
- CS Dep. Veterinary Pathobiol., Ohio State Univ., 1925 Coffey Road, Columbus, OH 43210, USA
- SO Infection and Immunity, (1994) Vol. 62, No. 9, pp. 3604-3607. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 24 Oct 1994
- Last Updated on STN: 25 Oct 1994
- AB Thirty-seven gnotobiotic piglets from seven litters were infected with either Helicobacter pylori N6 or urease-negative H. pylori N6ureG::Km, which contains an insertion in the ureG gene and produces inactive urease. To produce achlorhydria, piglets were treated throughout the experiment with omeprazole (5 mg intravenously every 12 h) and ranitidine (75 mg orally every 6 h). Treatment resulted in elevation of gastric pH to 7.0 +- 1.1 throughout the experiment. Control piglets were not treated and remained normochlorhydric. Strain N6 colonized well in both normal and achlorhydric piglets. All 10 piglets were colonized, and colonization ranged from 4.4 +- 1.5 log,, CFU/g of gastric mucosa in normochlorhydric piglets sacrificed after 2 days to 6.9 +- 0.5 log-10 CFU/g in normochlorhydric piglets sacrificed after 5 days. Strain N6ureG::Km did not colonize any of seven normochlorhydric piglets and was recovered only in low numbers (lt 100 CFU/g) from four of nine achlorhydric piglets. In the second experiment, piglets were coinoculated with both strains N6 and N6ureG::Km. Coinoculation did not affect colonization by urease-positive ***Urease*** - ***deficient*** N6ureG::Km was unable to colonize even in the presence of urease-positive ***bacteria*** . These results confirm that urease enzymatic activity (and not urease protein) is essential for colonization, that this effect is independent of diffusible products of urea metabolism, and that gastric pH protection is not a major role of urease in promoting colonization by H. pylori.
- AB. . . the second experiment, piglets were coinoculated with both strains N6 and N6ureG::Km. Coinoculation did not affect colonization by urease-positive N6. ***Urease*** ***deficient*** N6ureG::Km was unable to colonize even in the presence of urease-positive
 - ***bacteria*** . These results confirm that urease enzymatic activity (and not urease protein) is essential for colonization, that this effect is independent. . .

ORGN Classifier

Aerobic Helical or Vibrioid Gram-Negatives 06210 Super Taxa

Eubacteria; ***Bacteria*** ; Microorganisms

```
Organism Name
        aerobic helical or vibrioid gram-negative ***bacteria***
        Helicobacter pylori
     Taxa Notes
            ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
       Suidae
               85740
     Super Taxa
       Artiodactyla; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
       pig
     Taxa Notes
       Animals, Artiodactyls, . . .
L11 ANSWER 13 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on
     1993:333419 BIOSIS <<LOGINID::20080330>>
    PREV199345028144
DN
    An isogenic ***urease*** - ***deficient*** mutant of Helicobacter
    pylori colonizes gastric epithelial explants, but not germ-free piglets.
AU
     Eaton, K. A. [Reprint author]; Labigne, A. F.; Krakowka, S.
CS
   Dep. Vet. Pathobiol., Ohio State Univ., Columbus, OH, USA
   Gastroenterology, (1993) Vol. 104, No. 4 SUPPL., pp. A694.
SO
    Meeting Info.: 94th Annual Meeting of the American Gastroenterological
    Association. Boston, Massachusetts, USA. May 15-21, 1993.
    CODEN: GASTAB. ISSN: 0016-5085.
DT
    Conference; (Meeting)
T.A
   English
ED
   Entered STN: 16 Jul 1993
     Last Updated on STN: 31 Aug 1993
    An isogenic
                 ***urease*** - ***deficient*** mutant of Helicobacter
     pylori colonizes gastric epithelial explants, but not germ-free piglets.
ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives
     Super Taxa
        Eubacteria; ***Bacteria*** ; Microorganisms
     Organism Name
        aerobic helical or vibrioid gram-negative ***bacteria***
        Helicobacter pylori
     Taxa Notes
            ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
       Suidae 85740
     Super Taxa
        Artiodactyla: Mammalia: Vertebrata: Chordata: Animalia
     Organism Name
       Suidae
     Taxa Notes
       Animals, Artiodactyls,. . .
L11 ANSWER 14 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on
     SIN
AN
    1992:327557 BIOSIS <<LOGINID::20080330>>
DN
   PREV199294029398; BA94:29398
   CHARACTERIZATION OF HELICOBACTER-PYLORI UREASE MUTANTS.
TΙ
AU SEGAL E D [Reprint author]; SHON J; TOMPKINS L S
CS
    DEP MICROBIOL IMMUNOL, DIGESTIVE DISEASES CENTER, STANFORD UNIV, STANFORD,
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CALIF 94305, USA Infection and Immunity, (1992) Vol. 60, No. 5, pp. 1883-1889. SO CODEN: INFIBR. ISSN: 0019-9567. DT Article FS T.A ENGLISH ED Entered STN: 11 Jul 1992 Last Updated on STN: 11 Jul 1992 The association between Helicobacter pylori, gastritis, and peptic ulcer ΔR is well established, and the association of infection with gastric cancer has been noted in several developing countries. However, the pathogenic mechanism(s) leading to disease states has not been elucidated. The H. pylori urease is thought to be a determinant of pathogenicity, since the enzyme is produced by all H. pylori clinical isolates. Evidence indicates that some H. pylori strains are more cytotoxic than others, with a correlation between the activity of the urease and the presence of a vacuolating cytotoxin having been made. However, the number of cytotoxins remains unknown at this time. The relationship between the urease and cytotoxicity has previously been examined with chemical inhibitors. To examine the role of the urease and its relationship to cytotoxicity, ***urease*** - ***deficient*** mutants were produced following ethyl methanesulfonate mutagenesis of H. pylori 87A300. Two mutants (the ure1 and ure5 mutants) which were entirely deficient in urease activity (Ure-) were selected. Characterization of the isolates at the protein level showed that the urease subunits lacked the ability to complex and form the active urease enzyme. The urel mutant was shown to be sensitive to the effects of low pH in vitro and exhibited no cytotoxicity to eucaryotic cells, whereas the parental strain (Ure+) produced a cytotoxic effect in the presence of urea. Interaction between the H. pylori Ure+ and Ureproduced a cytotoxic effect in the presence of urea. Interaction between the H. pylori Ure+ and Ure- strains and Caco-2 cells appeared to be similar in that both ***bacterial*** types elicited pedestal formation and actin condensation. These results indicate that the H. pylori ureas may have many functions, among them (i) protecting H. pylori against the acidic environment of the stomach, (ii) acting as a cytotoxin, with human gastric cells especially susceptible to its activity, and (iii) disrupting cell tight junctions in such a manner than the cells remain viable but an ionic flow between the cells occurs. AB. . . cytotoxicity has previously been examined with chemical inhibitors. To examine the role of the urease and its relationship to cytotoxicity, ***urease*** - ***deficient*** mutants were produced following ethyl methanesulfonate mutagenesis of H. pylori 87A300. Two mutants (the ure1 and ure5 mutants) which were. . . urea. Interaction between the H. pylori Ure+ and Ure- strains and Caco-2 cells appeared to be similar in that both ***bacterial*** types elicited pedestal formation and actin condensation. These results indicate that the H. pylori ureas may have

many functions, among. .

ORGN Classifier
Aerobic Helical or Vibrioid Gram-Negatives 06210
Super Taxa
Eubacteria; ***Bacteria*** ; Microorganisms
Taxa Notes

Bacteria , Eubacteria, Microorganisms
ORGN Classifier
Vertebrata 85150
Super Taxa
Chordata; Animalia

Taxa Notes
Animals, Chordates, Nonhuman Vertebrates, Vertebrates

L11 ANSWER 15 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on AN 1992:41708 BIOSIS <<LOGINID::20080330>> DN PREV199242017858; BR42:17858 TT CONSTRUCTION OF ***UREASE*** ***DEFICIENT*** MUTANTS OF HELICOBACTER-PYLORI BY ALLELIC EXCHANGE. AII FERRERO R [Reprint author]; CUSSAC V; COURCOUX P; LABIGNE A CS UNITE ENTEROBACTERIES, INSERM U199, INST PASTEUR, 75724 PARIS CEDEX 15, FR Microbial Ecology in Health and Disease, (1991) Vol. 4, No. SPEC. ISSUE, SO pp. S136. Meeting Info.: VITH INTERNATIONAL WORKSHOP ON CAMPYLOBACTER HELICOBACTER AND RELATED ORGANISMS, SYDNEY, NEW SOUTH WALES, AUSTRALIA, OCTOBER 7-10, 1991. MICROB ECOL HEALTH DIS. ISSN: 0891-060X. DТ Conference; (Meeting) FS BR LA ENGLISH Entered STN: 7 Jan 1992 ED Last Updated on STN: 8 Jan 1992 CONSTRUCTION OF ***UREASE*** ***DEFICIENT*** MUTANTS OF HELICOBACTER-PYLORI BY ALLELIC EXCHANGE. ORGN Classifier Aerobic Helical or Vibrioid Gram-Negatives 06210 Super Taxa Eubacteria; ***Bacteria*** ; Microorganisms Taxa Notes ***Bacteria*** , Eubacteria, Microorganisms ORGN Classifier Enterobacteriaceae 06702 Facultatively Anaerobic Gram-Negative Rods; Eubacteria; ***Bacteria*** ; Microorganisms Taxa Notes ***Bacteria*** , Eubacteria, Microorganisms L11 ANSWER 16 OF 20 CABA COPYRIGHT 2008 CABI on STN AN 95:23391 CABA <<LOGINID::20080330>> DN 19941908449 TI Hydrogenase and urease in cyanobacterial photosynthesis and nitrogen fixation Ewart, G. D.; Mackerras, A. H.; Smith, G. D.; Kashyap, A. K. [EDITOR]; Kumar, H. D. [EDITOR] 22 Department of Biochemistry, Fculty of Science, Australian National University, Canberra, ACT 2601, Australia. Recent advances in phycology, (1994) pp. 21-30. 26 ref. Publisher: Rastogi Publications. Meerut ISBN: 81-85711-05-4 CY India DT Miscellaneous LA English

AB In the cyanobacterium Anabaena cylindrica both hydrogenase and urease activities are dependent on the presence of Ni in the growth medium. In

ED

Entered STN: 1 Feb 1995 Last Updated on STN: 1 Feb 1995 cyanobacteria there are two forms of hydrogenase: soluble and membrane bound. Electrophoretic analysis showed that the enzyme is a dimer consisting of 2 subunits. Tritium exchange and reductive hydrogenase activities could be differentially inhibited. Growth of cells in the absence of Ni produced hydrogenase and **urease** - ***deficient*** cells. The exponential growth rate of nitrogen-fixing cells in A. cylindrica was not inhibited by the absence of Ni. Growth of A. cylindrica was dependent on Ni when non-nitrogen-fixing cells were used to reinitiate nitrogen-fixing growth. Nickel-deficient cells showed a pronounced growth lag which was associated with loss of pigment, delayed nitrogenase synthesis, and cyanophycin accumulation. These observations suggested a role for Ni in nitrogen metabolism in addition to that as a cofactor for urease.

AB . . exchange and reductive hydrogenase activities could be differentially inhibited. Growth of cells in the absence of Ni produced hydrogenase and ***urease*** - ***deficient*** cells. The exponential growth rate of nitrogen-fixing cells in A. cylindrica was not inhibited by the absence of Ni. Growth.

ORGN ***bacteria*** ; Cyanobacteria

- L11 ANSWER 17 OF 20 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2006:380705 CAPLUS <<LOGINID::20080330>>
- DN 144:410795
- TI Recombinant ***Mycobacterium*** BCG adjuvant in vaccination
- IN Laeufer, Albrecht; Eisele, Bernd; Grode, Leander
- PA Vakzine Projekt Management G.m.b.H., Germany
- SO Eur. Pat. Appl., 17 pp.
- CODEN: EPXXDW
- DT Patent
- LA English
- DAN CHE 1

FAN.CNT 1 PATENT NO.				KIND		DATE		APPLICATION NO.					Di	ATE					
PI	EP	1649	869			A1		2006	0426		EP 2	004-	2509	6		21	0041	021	
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,	
			ΙE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	HU,	PL,	SK,	į
	AU	2005	2989	76		A1		2006	0504		AU 2	005-	2989	76		21	0051	016	
	CA	2584	321			A1		2006	0504		CA 2	005-	2584	321		21	0051	016	
	WO	2006	0454	68		A1		2006	0504		WO 2	005-	EP11	127		21	0051	016	
		W:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	ΒZ,	CA,	CH,	
			CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	ΓI,	GB,	GD,	
			GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KΕ,	KG,	KM,	KΡ,	KR,	ΚZ,	
								LU,											
			NΑ,	NG,	ΝI,	NO,	ΝZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	
			SK,	SL,	SM,	SY,	ΤJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UΖ,	VC,	VN,	
					ZM,														
		RW:						CZ,											
								MC,											
								GN,											
								NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	ΑZ,	BY,	
						RU,													
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		1010																	
		2007																	
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HR

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RR 2007068398 A 20070629 KR 2007-709076 20070420 PRAI EP 2004-25096 A 20041021 WO 2005-EP11127 W 20051016
   The authors disclose the use of ***urease*** - ***deficient***
       ***Mycobacterium*** BCG expressing listeriolysin as an adjuvant in
     vaccination. In one example, a tumor vaccine comprises a allogeneic
     prostate carcinoma cell line, transgenic for interferon-.gamma. and
     interleukin-2, in combination with the foregoing ***bacterial*** cell
     adiuvant.
             THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 6
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
    Recombinant ***Mycobacterium*** BCG adjuvant in vaccination
AB
     The authors disclose the use of ***urease*** - ***deficient***
       ***Mycobacterium*** BCG expressing listeriolysin as an adjuvant in
     vaccination. In one example, a tumor vaccine comprises a allogeneic
     prostate carcinoma cell line, transgenic for interferon-.gamma. and
     interleukin-2, in combination with the foregoing ***bacterial*** cell
     adjuvant.
ST
      ***Mycobacterium*** cytolysin adjuvant vaccine
ΙT
    Vaccines
        (antimalarial; ***urease*** - ***deficient***
          ***Mycobacterium*** BCG expressing listeriolysin as adjuvant for)
ΙT
     Antigens
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (autoantigens, microbial; ***urease*** - ***deficient***
          ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
        vaccination against)
    Prostate gland, neoplasm
TT
        (carcinoma; ***urease*** - ***deficient*** ***Mycobacterium***
        BCG expressing listeriolysin as vaccine adjuvant for
        cytokine-transgenic cell immunogens)
TТ
   Intestine, neoplasm
        (colon, carcinoma; ***urease*** - ***deficient***
          ***Mycobacterium*** BCG expressing listeriolysin as vaccine adjuvant
        for cytokine-transgenic cell immunogens)
ΙT
    Carcinoma
                ***urease*** - ***deficient***
                                                    ***Mvcobacterium***
        (colon;
        BCG expressing listeriolysin as vaccine adjuvant for
        cvtokine-transgenic cell immunogens)
ΙT
    Carcinoma
        (head and neck squamous cell carcinoma; ***urease***
          ***deficient*** ***Mycobacterium*** BCG expressing listeriolysin
        as vaccine adjuvant for cytokine-transgenic cell immunogens)
    Cell adhesion molecules
     Interleukin 12
     Interleukin 2
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (in combination with ***urease*** - ***deficient***
          ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
       vaccination)
    Hemolvsins
     RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use): BIOL (Biological study): USES (Uses)
        (listeriolysins 0; ***urease*** - ***deficient***
          ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
       vaccination)
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IT Antigens
    Tumor antigens
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (microbial; ***urease*** - ***deficient*** ***Mycobacterium***
       BCG expressing listeriolysin as adjuvant in vaccination against)
TT
   Lung, neoplasm
        (non-small-cell carcinoma;
                                  ***urease*** - ***deficient***
         ***Mycobacterium*** BCG expressing listeriolysin as vaccine adjuvant
       for cytokine-transgenic cell immunogens)
TT
    Lysosome
        (phagolysosome; ***urease*** - ***deficient***
         ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
       vaccination in relation to)
ΙT
    Carcinoma
        (prostatic; ***urease*** - ***deficient*** ***Mycobacterium***
       BCG expressing listeriolysin as vaccine adjuvant for
       cvtokine-transgenic cell immunogens)
ΙT
    Carcinoma
        (pulmonary non-small-cell; ***urease*** - ***deficient***
         ***Mycobacterium*** BCG expressing listeriolysin as vaccine adjuvant
       for cytokine-transgenic cell immunogens)
IΤ
    Kidney, neoplasm
                              ***urease*** - ***deficient***
       (renal cell carcinoma;
         ***Mycobacterium*** BCG expressing listeriolysin as vaccine adjuvant
       for cytokine-transgenic cell immunogens)
TΤ
    Carcinoma
        (renal cell: ***urease*** - ***deficient*** ***Mycobacterium***
       BCG expressing listeriolysin as vaccine adjuvant for
       cytokine-transgenic cell immunogens)
TТ
    Head and Neck, neoplasm
                                  ***urease*** - ***deficient***
        (squamous cell carcinoma;
         ***Mycobacterium*** BCG expressing listeriolysin as vaccine adjuvant
       for cytokine-transgenic cell immunogens)
ΙT
    Vaccines
        (tumor:
               ***urease*** - ***deficient***
                                                    ***Mycobacterium***
       BCG expressing listeriolysin as adjuvant for)
ΙT
    MSP-1 (protein)
    RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
       ( ***urease*** - ***deficient***
                                             ***Mycobacterium***
                                                                    BCG
       expressing listeriolysin as adjuvant for)
    Plasmodium falciparum
       ( ***urease*** - ***deficient***
                                            ***Mvcobacterium***
       expressing listeriolysin as adjuvant for merozoite surface protein of)
    Malaria
       ( ***urease*** - ***deficient***
                                             ***Mycobacterium*** BCG
       expressing listeriolysin as adjuvant for vaccination against)
ΙT
    Human
        ***Mycobacterium***
                            BCG
        ***Mvcobacterium***
                             ***bovis***
        ( ***urease*** - ***deficient***
                                              ***Mycobacterium*** BCG
       expressing listeriolysin as adjuvant in vaccination)
    Antigen-presenting cell
    Brain, neoplasm
    Dendritic cell
    Mammary gland, neoplasm
    Melanoma
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Neoplasm
       ( ***urease*** - ***deficient*** ***Mycobacterium*** BCG
       expressing listeriolysin as vaccine adjuvant for cytokine-transgenic
       cell immunogens)
ΙT
    Antimalarials
    Antitumor agents
        (vaccines: ***urease*** - ***deficient***
                                                        ***Mycobacterium***
       BCG expressing listeriolysin as adjuvant for)
    Interferons
    RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
                                      ***urease*** - ***deficient***
        (.gamma.; in combination with
         ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
       vaccination)
ΤT
    884349-82-0
    RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (amino acid sequence; ***urease*** - ***deficient***
         ***Mvcobacterium***
                             BCG expressing listeriolysin as adjuvant in
       vaccination)
    9002-13-5D, Urease, subunit C
ΙT
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (deficiency; ***urease*** - ***deficient***
                                                          ***Mvcobacterium***
       BCG expressing listeriolysin as adjuvant in vaccination)
    884349-81-9, DNA (Listeria monocytogenes gene hyl)
    RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (nucleotide sequence; ***urease*** - ***deficient***
         ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
       vaccination)
L11 ANSWER 18 OF 20 CAPLUS COPYRIGHT 2008 ACS on STN
AN
    1997:498189 CAPLUS <<LOGINID::20080330>>
DN
    127:188074
ΤI
    Interactions of a catalase- and an urease-negative mutant of Helicobacter
    pylori with polymorphonuclear granulocytes
ΑU
    Marxer, Martin; Farzam, Fardad; Spiegelhalder, Christiane; Kersten,
    Astrid; Odenbreit, Stefan; Haas, Rainer; Kist, Manfred
    Inst. fur Med. Mikrobiologie und Hygiene, Freiburg, 79104, Germany
SO
    Campylobacters, Helicobacters, and Related Organisms, [Proceedings of the
    International Workshop on Campylobacters, Helicobacters, and Related
    Organisms], 8th, Winchester, UK, July 10-13, 1995 (1996), Meeting Date
    1995, 701-705. Editor(s): Newell, Diane G.; Ketley, Julian M.; Feldman,
    Roger A. Publisher: Plenum, New York, N. Y.
    CODEN: 64TNAY
DT
    Conference
LA English
```

factors of H. pylori, isogenic catalase- or ***urease*** -***deficient*** mutant strains, constructed by transposon mutagenesis, were compared with the corresponding wild-type strain 69A with respect to their interactions with polymorphonuclear nucleophiles (PMNs), including sensitivity towards killing by PMNs, strength of the oxidative burst, and electron microscopic studies. The results from the the catalase-neg. mutant indicated that although catalase is able to scavenge hydrogen peroxide, it does not protect the ***bacteria*** efficiently from

To examine whether or not catalase and urease play a role as virulence

CS

AB

PMN-induced killing. In the case of the urease-neg. mutant, the phagocytic oxidative burst in the presence of the mutant was not significantly increased compared to that induced by the wild type, thus suggesting that non-oxygen-mediated killing mechanisms of the PMNs are responsible form the more efficient ***bracese*** - ***deficient*** mutant.

AB To examine whether or not catalase and urease play a role as virulence factors of H. pylori, isogenic catalase- or ***urease*** -

deficient mutant strains, constructed by transposon mutagenesis, were compared with the corresponding wild-type strain 69A with respect to their interactions with. . . from the the catalase-neg. mutant indicated that although catalase is able to scavenge hydrogen peroxide, it does not protect the ***bacteria*** efficiently from PMN-induced killing. In the case of the urease-neg. mutant, the phagocytic oxidative burst in the presence of the. . induced by the wild type, thus suggesting that non-oxygen-mediated killing mechanisms of the PMNs are responsible form the more efficient ***urease*** - ***deficient*** mutant.

- L11 ANSWER 19 OF 20 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 1992:172374 CAPLUS <<LOGINID::20080330>>
- DN 116:172374
- TI Selection of L-lysine-producing strain Aul11-2
- AU Su, Lingming; Xu, Suowei; Fu, Yinghua; Wang, Xingzhen; Tang, Shanghua; Shao, Guoliang
- CS Shanghai Inst. Ind. Microbiol., Shanghai, Peop. Rep. China
- SO Gongye Weishengwu (1991), 21(6), 12-16
- CODEN: GOWEEK; ISSN: 1001-6678
- DT Journal
- LA Chinese
- AB ***Bacteria*** strain All1 was a good producer of lysine, but was
 urease ***deficient*** , and so the pH in the process of
 fermn.

could not be controlled with urea. After the mutation with MNNG and screening with urea as nitrogen source, an urease revertant strain Au11-2 was obtained. The lysine productivity and the conversion ratio to the glucose of the urease revertant Au111-2 increased by 25% and 15% than that of strain Au11.

AB ***Bacteria*** strain All1 was a good producer of lysine, but was ***urease*** ***deficient*** , and so the pH in the process of

could not be controlled with urea. After the mutation with MNNG. . . ST lysine fermn ***bacteria*** urease

- IT ***Bacteria***
- (lysine formation by, urease mutation effect on)
- IT Fermentation
 - (lysine, with ***bacteria*** , urease mutation effect on)
- IT 56-87-1, L-Lysine, biological studies
- RL: FORM (Formation, nonpreparative)
- (formation of, by ***bacteria*** , urease mutation effect on)
- IT 9002-13-5, Urease
- RL: BIOL (Biological study)
- (of ***bacteria*** , lysine formation in relation to)
- L11 ANSWER 20 OF 20 MEDLINE on STN
- AN 2007476473 MEDLINE <<LOGINID::20080330>>
- DN PubMed ID: 17519853

- TI [Strategies for the development of new ***tuberculosis*** vaccines]. Strategie per lo sviluppo di nuovi vaccini antitubercolari.
- AU Fattorini L
- CS Dipartimento di Malattie Infettive, Parassitarie e Immunomediate, Istituto Superiore di Sanita, Roma, Italy.. lanfranco.fattorini@iss.it
- SO Minerva medica, (2007 Apr) Vol. 98, No. 2, pp. 109-19. Ref: 47 Journal code: 0400732. ISSN: 0026-4806.
- CY Italy
- DT (ENGLISH ABSTRACT)

 Journal; Article; (JOURNAL ARTICLE)
 - General Review; (REVIEW)
- LA Italian
- FS Priority Journals
- EM 200708
- ED Entered STN: 16 Aug 2007
- Last Updated on STN: 17 Aug 2007
- Entered Medline: 16 Aug 2007
- AB ***Tuberculosis*** remains a substantial global health problem causing 2 million deaths, and an estimated 8 to 10 million new infections a year. The efficacy of the ***Mycobacterium*** ***bovis*** Bacillus Calmette-Guerin (BCG), the only available antituberculosis vaccine, is variable (0-80%), especially in ***tuberculosis*** -endemic countries. Over the past decade there has been a resurgence of interest in the development of new ***tuberculosis*** vaccines and some of the most promising are now entering into early clinical trials, based on two different strategies. The first is to use whole **mycobacteria*** to replace BCG (priming vaccines), either by developing a recombinant strain of BCG or an attenuated strain of ***Mycobacterium***
 - ***tuberculosis*** . To date, two recombinant strains of BCG, one overexpressing antigen 85B (rBCG-85B) and the other, a ***urease*** ***deficient*** BCG mutant which expresses the listeriolygin O gene

from

- Listeria monocytogenes (rBGG::DeltaureC-hly+), entered into clinical trials. The second approach is to develop subunit vaccines (recombinant proteins and viral vectors, and DNA vaccines) expressing immunodominant antigen/s from M. ***tuberculosis*** able to augmenting BCG protection (booster vaccines). At the moment, three major vaccines, namely a recombinant modified vaccinia virus Ankara expressing antigen 85A (WNA95A), a fusion protein of SEAT6 and 85B (Hybrid 1), and another fusion protein comprising the 32 and 39 Kda proteins (72f) entered into clinical trials.
- TI [Strategies for the development of new ***tuberculosis*** vaccines].

 Strategie per lo sviluppo di nuovi vaccini antitubercolari.
- AB ***Tuberculosis*** remains a substantial global health problem causing 2 million deaths, and an estimated 8 to 10 million new infections a year. The efficacy of the ***Mycobacterium*** ***bovis*** Bacillus Calmette-Guerin (BCG), the only available antituberculosis vaccine, is variable (0-80%), especially in ***tuberculosis*** -endemic countries. Over the past decade there has been a resurgence of interest in the development of new ***tuberculosis*** vaccines and some of the most promising are now entering into early clinical trials, based on two different strategies. The first is to use whole ***mycobacteria*** to replace BCG (priming vaccines), either by developing a recombinant strain of BCG or an attenuated strain of ***Mycobacterium***
 - ***tuberculosis*** . To date, two recombinant strains of BCG, one overexpressing antiqen 85B (rBCG-85B) and the other, a ***urease*** -

```
***deficient*** BCG mutant which expresses the listeriolysin 0 gene

Listeria monocytogenes (rBCG::DeltaureC-hly+), entered into clinical

trials. The second approach is to develop subunit vaccines (recombinant
proteins and viral vectors, and DNA vaccines) expressing immunodominant
antigen/s from M. ***tuberculosis*** able to augmenting BCG protection
(booster vaccines). At the moment, three major vaccines, namely a
recombinant modified vaccinia virus Ankara.

CI Immunization, Secondary: MT, methods

****Nycobacterium bovis: IM, immunology***

****Mycobacterium tuberculosis: IM, immunology***

****Tuberculosis Vaccines: IM, immunology

Vaccines, Synthetic: IM, immunology

CN 0 (***Tuberculosis*** Vaccines); 0 (Vaccines, Synthetic)
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